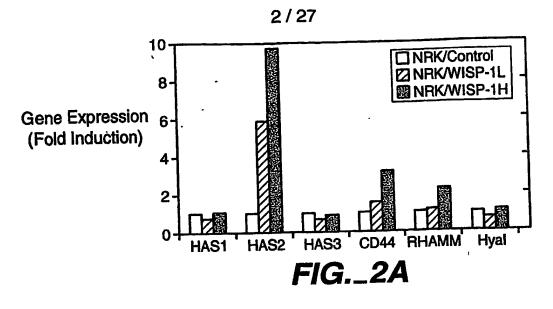


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Frequency

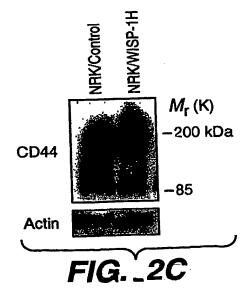
NRK/Control

NRK/WISP-1H

1.0 10.0 100.0

FIG._2B

CD44 Expression (Fluorescence Intensity)



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3/27 FIG._3C FIG._3B FIG._3A NRKWISP-1L NRK/Contre NRKWISP NRK/Control FIG._3E FIG._3D 160 120 Migration Distance 80 (µm) 40 FIG._3F NRK/Control NRK/WISP-1H 600 400 Migration Area (μm²) 200 FIG. 3G 'NRK/WISP-1H NRK/Control SUBSTITUTE SHEET (RULE 26)



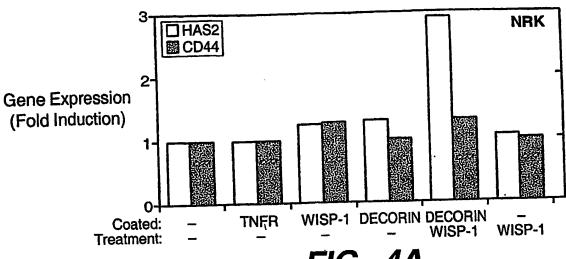


FIG._4A

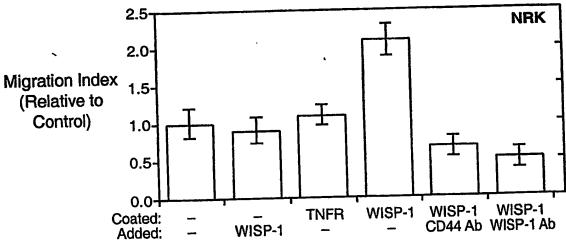


FIG._4B

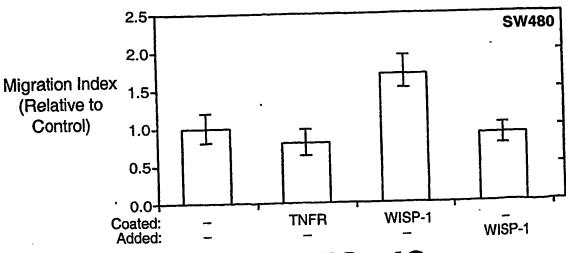
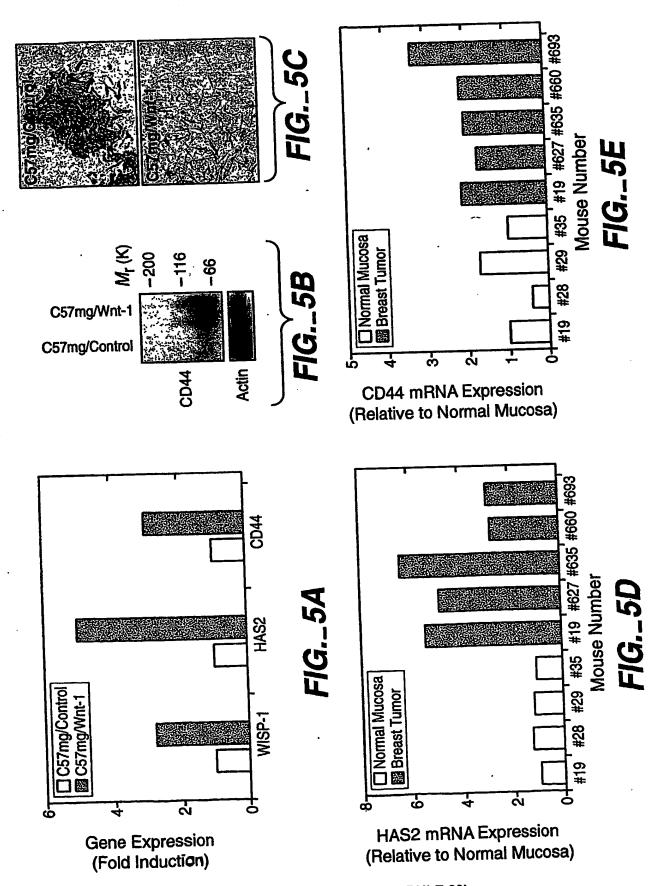
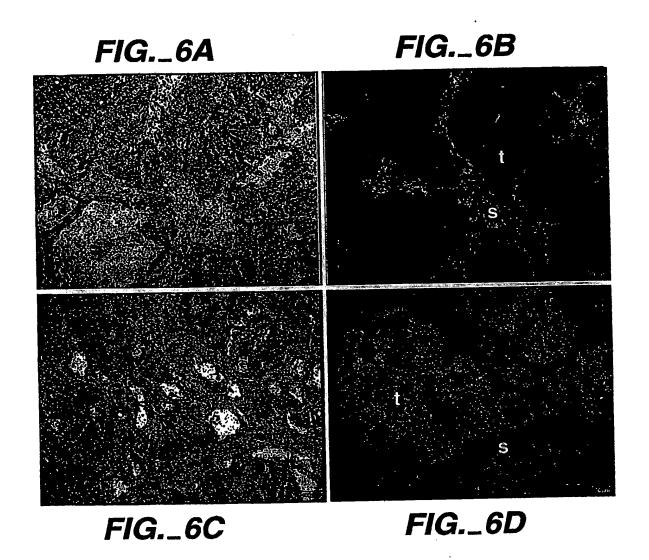
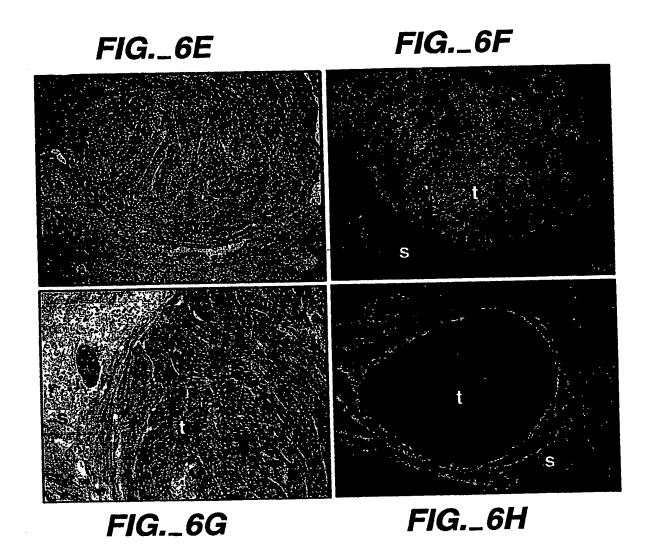
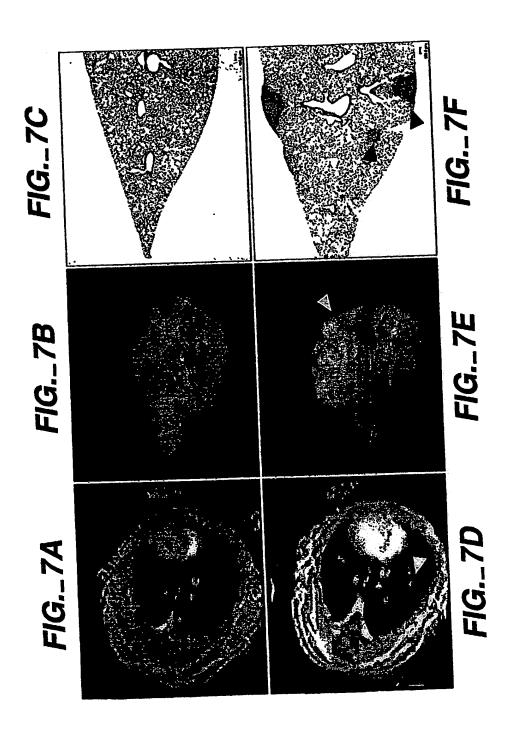


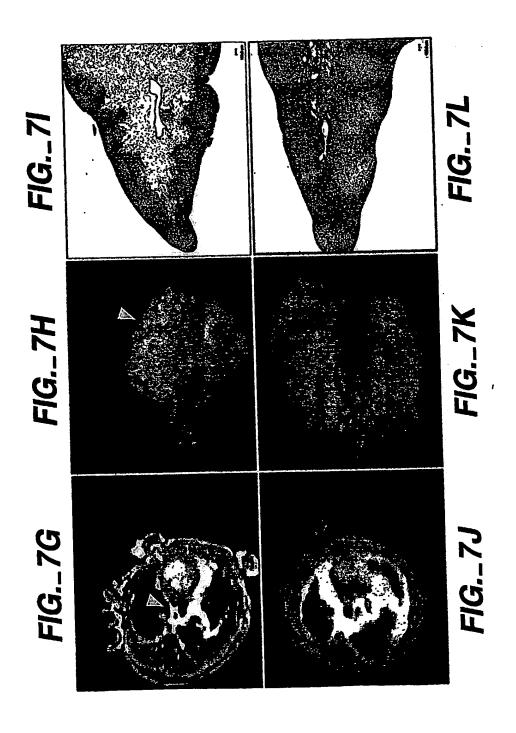
FIG._4C



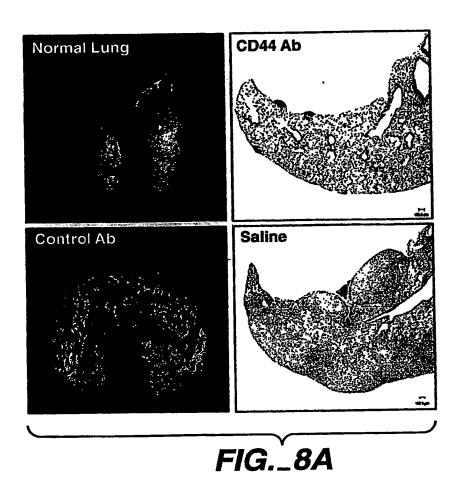


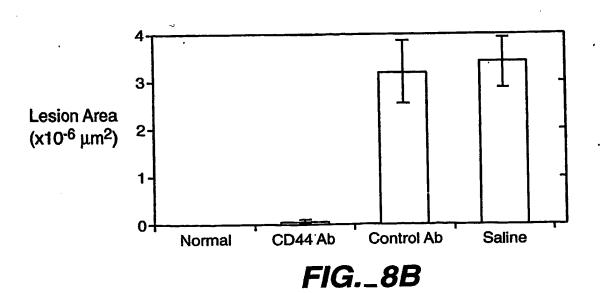






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TCCACTCGGA

CCTGGGGGTC

TGAGGTCGAG CATGGACTTT ACTCCAGCTC TCGAGGGGC TCTCCACCAG CCTAGGAGAC CCGACGAGCC AGCTACGGAC ACGTGACTG CAGGTCCGTA CTCCACCAAG GGAICCTCTG GGCTGCTCGG ICGAIGCCTG IGCCACTGAC GTCCAGGCAI GAGGIGGITC GTACCTGAAA Ω Σ 101 CHGCCCTGGA CGCTGGCAGC AGTGACAGCA GCAGCGGCCA GCACCGTCCT GGCCACGGCC CTCTCCCAG CCCCTACGAC GAGAGAGGTC Д ຜ ... CGTCGGCGGT CGTGGCAGGA CCGGTGCCGG E٠ æ > E4 1 CCCACGCGTC CGCTGGGCCC AGCTCCCCCG AGAGGTGGTC e e TCACTGTCGT æ > GCGACCGTCG GGGTGCGCAG GCGACCCGGG æ L GACGGGACCT

CCCCAGTCGG AGTAGTGTCT GGGGTCAGCC TCATCACAGA ß ა ე CTGCCCGCTG GACGGGCGAC H U TGCCCGCCAT CCCCACCCCG ACGGGCGGTA GGGGTGGGGC œ Д ф Д Д CGCCCCCAAT TCTGCAAGTG GCCATGTGAG CGGTACACTC Ŋ ρ, GCGGGGTTA AGACGTTCAC × ပ O Д æ 201 CACTGGAGGA CACCTCCTCA GTGGAGGAGT ഗ ល H GTGACCTCCT 33

CCCCACCGGG GCCTCTACTG TGACTACAGC GGGGTGGCCC CGGAGATGAC ACTGATGTCG × ធ U œ H Д CATCTGTGAC GTAGACACTG Ω U Н GCCTCCGACG TGGCTGTGAG TGCTGTAAGA TGTGCGCTCA GCAGCTTGGG GACAACTGCA CGGAGGCTGC Ø H CTGTTGACGT ပ Z CGTCGAACCC U O ACCGACACTC ACGACATTCT ACACGCGAGT O 4 U × U ഠ ပ Ö 301 22

AGGAAGGTCG CAACGGCCAG TCCTTCCAGC ĵ24 ຜ GITGCCGGTC O z CTGGATGGGG TGCGCTACAA GACCTACCCC ACGCGATGTT × œ > Ö U D GGGCTGCGTC CCCGACGCAG > ບ Ö GGGGACCGCC CGAGGTACGC AATAGGAGTG TGTGCACAGG TGGTCGGTGT ACCAGCCACA > Ö > TTATCCTCAC ACACGTGTCC Ø ď ບ > U Н GCTCCATGCG CCCCTGGCGG G 105 401

CCACTGTGCC TCCGAGTGCG CCCCCGCGT CTCTGGTGCC CCCACCCGCG GGGTGGGCGC GAGACCACGG GGTGACACGG AGGCTCACGC GGGGGGCGCA × ۵, م > ĸ ပ ъ П GGGCTGCACA TECCECECCA CCCGACGTGT H U O GTACAACTGC ACGTGCATCG ACGGCGCGT A V Ö CATGTTGACG TGCACGTAGC н ပ U z × CTAACTGCAA GATTGACGTT 501 139

TGTGAGGACG ACGCCAAGAG GCCACGCAAG ACCGCACCCC GTGACACAGG AGCCTTCGAT TCGGAAGCTA Ω CACTGTGTCC E4 TGGCGTGGGG Ç. 4 CGGTGCGTTC × 24 Д TGCGGTTCTC œ × 4 CGTCACCCAT ACACTCCTGC Ω Ω M ပ GCGCGTGAGC ATACCTGGCC ACTGCTGTGA GCAGTGGGTA > 3 OI TGACGACACT ဂ ங် ပ TATGGACCGG Ö CGCGCACTCG 601 172

GGACCCCCAG AGGTGAGCCT Ö ᆸ AACTGCATAG CCTACAAG CCCCTGGAGC CCTTGCTCCA CCAGCTGCGG GGAACGAGGT GGTCGACGCC Ö ပ ß H ςΩ U Д GGATGTTC GGGGACCTCG ß 3 ß H × TTGACGTATC н ပ AGGTGGAGGC ATGGCACAGG TACCGTGTCC ρá ;;; 3 TCCACCTCCG 臼 > CGACACCCAC GCTGTGGGTG 4 701

TGCTGGCCTG AGCAAGAGA CCGCCTCTGC AACTTGCGGC CATGCGATGT GGACATCCAT ACACTCATTA AGGCAGGGAA GGCGGAGACG TTGAACGCCG GTACGCTACA CCTGTAGGTA TGTGAGTAAT TCCGTCCCTT J Н Α Ω о Д œ r O p4 ACGACCGGAC TCGTTCTCTC Ŋ 臼 Ø 3 ပ AGAGGTTACA ATTGCGGGTC TAACGCCCAG a 4 TCTCCAATGT 801 239

AATTCAAAGA GAAGTGGGGA TGTGACACTT CCCATGTCTA ATCCAAACAG GGTCAGTCTT TATTTTAAAC TATTTGTAAG GACAACTACC CTTTTCGGGG CAGITAATAC TCCAGAGACA GGGAAAGGIC AGCCCATITC AGAAGGACCA AITGACICIC ACACIGAAIC AGCIGCIGAC TGGCAGGCT ITGGGCAGIT GCACCTCAAG GTCATCAAAC ATTIGCCAAG IGAGTIGAAT AGTIGCTIAA TITITGATTIT TAATGGAAAG TIGTATCCAT TAACCTGGGC ATTGTTGAGG CGTGGAGTTC CAGTAGTTTG TAAACGGTTC ACTCAACTTA TCAACGAATT AAAACTAAAA ATTACCTTTC AACATAGGTA ATTGGACCCG TAACAACTCC THAAGITITCT CITCACCCCT ACACIGIGAA GGGTACAGAI TAGGITIGIC CCAGICAGAA AIAAAAITIG AIAAACAIIC CIGITGAIGG GAAAAGCCCC CTTGTTATCT TAFTAGATAA TAAACCTCGG ACGGTTCTCC ATGACATTAC CCATTAAGAC TGCAGTCGCG TGGTTTTGAT AGGACTAAGG TTTATACATA GAACAATAGA ATAATCTATT ATTIGGAGCC TGCCAAGAGG TACTGTAATG GGTAATTCTG ACGTCAGCGC ACCAAAACTA TCCTGATTCC AAATATGTAT TTCTTTACCT TAGTCCATCT GAAAATTATA GTGATTAAAG AAGAAATCTA CGGTTTGGTG TTCTGAGAAA CCCAGGTAAG TCTACTTATC TACCTTAAAC GATITICTITIT TACGGACAGA GATCGACAAG ACCTGATGTG GGTTCGGACT AGGTCGGAAA GGTTCAGTGA TCTTCAGGAC GACCTAGAAC GGATTTAGGG ATGGAATTTG CITGGCCTCC ATTICTGTCT CTAACCATTC AAATGACGCC TGATGGTGCT GCTCAGGCCC ATGCTATGAG TITTCTCCTT GATATCATTC AGCATCTACT GAACCGGAGG TAAAGACAGA GATTGGTAAG TTTACTGCGG ACTACCACGA CGAGTCCGGG TACGATACTC AAAAGAGGAA CTATAGTAAG TCGTAGATGA CTGGATCTTG CCTAAATCCC GACTIGGAAT CCTACCCIGA CITCTCAGAA AITGCCAACT AGGCAGGCAC TAACGGTTGA TCCGTCCGTG 1201 AARTCTTGGG TCTTGGGGAC TAACCCAATG CCTGTGAAGC AGTCAGCCCT TATGGCCAAT AACTTTTCAC CAATGAGCCT TAGTTACCCT GATCTGGACC TITAGAACCC AGAACCCCTG AFTGGGTFAC GGACACTTCG TCAGTCGGGA AFACCGGTTA TTGAAAAGTG GFTACTCGGA ATCAATGGGA CTAGACCTGG GATACCTAAT CCTGATGGGC TIGGCTTCTC CCGCCAGGTC CTATGGATTA CGACACATGG TCGGTCTCCG TAGGTACTTG AAGTGTGAAC GCCCGACGTA GTCGTGTGCG AGGATAGTTG GGTTCATGAC ACCTCAAACG 901 GAAGTGTCTG GCTGTGTACC AGCCAGAGGC ATCCATGAAC TTCACACTTG CGGGCTGCAT CAGCACACGC TCCTATCAAC CCAAGTACTG TGGAGTTTGC AAGAAATGGA ATCAGGTAGA CTTTTAATAT CACTAATTTC TTCTTTAGAT GCCAAACCAC AAGACTCTTT GGGTCCATTC AGATGAATAG GGCGGTCCAG > 0 HAH œ ATGCCTGTCT CTAGCTGTTC TGGACTACAC CCAAGCCTGA TCCAGCCTTT CCAAGTCACT AGAAGTCCTG CTGAACCTTA GGATGGGACT GAAGAGTCTT D L E S Y P D F S E GGACTACCCG AACCGAAGAG Ö Ω A 1001 ATGGACAATA GGTGCTGCAT CCCCTACAAG TCTAAGACTA TCGACGTGTC CTTCCAGTGT ပ O [z4 TACGGACGAA GACATTGGAC TCGACATCCT TAGGGTTACT GTAGAAACGA A C F C N L S C R N P N D I F A 1101 ATGCCTGCTT CTGTAACCTG AGCTGTAGGA ATCCCAATGA CATCTTTGCT TACCTGTTAT CCACGACGTA GGGGATGTTC AGATTCTGAT AGCTGCACAG L M E S Ø × 臼 > Д, Д ۲ ۸ ပ ပ CTTCACAGAC ပ D Z 1801 1701 1601 1501 1401 1301 272 SUBSTITUTE SHEET (RULE 26)

GICAATTATG AGGICTCIGT CCCTTICCAG TCGGGTAAAG TCTTCCTGGT TAACTGAAG TGTGACTTAG TCGACGACTG ACCGTCCAA AACCCGTCAA

1901

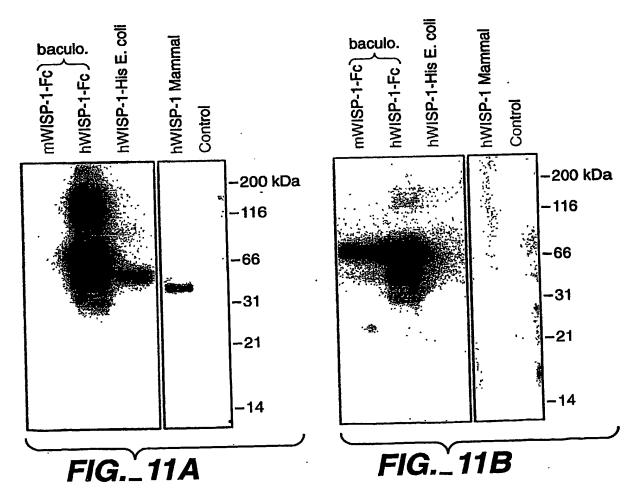
FIG._9C

AAGTCTAGCT TTAACCTGAC AGAAATATTG GGTATAAAAG GGGGACAAAA ATCTCGAAGG TTTACACAGT CTTATCCTTT TGTAACGTTA TTTACCGAAC CTGTGCCAGA CATTGCTCTC AGTGCTTTGC ATGTATTAGC TCACTGAATC TTCACGACAA TGTTGAGAAG 2601 TYCCCATTAT TATTTCTGTT CTTACAAATG TGAAACGGAA GCTCATAGAG GTGAGAAAAC TCAACCAGAG TCACCCAGTT GGTGACTGGG AAAGTTAGGA AAGGCTAATA ATAAAGACAA GAATGTTTAC ACTTTGCCTT CGAGTATCTC CACTCTTTTG AGTTGGTCTC AGTGGGTCAA CCACTGACCC TTTCAATCCT TTCAGATCGA AATTGGACTG TCTTTATAAC CCATATTTTC CCCCTGTTTT TAGAGCTTCC AAATGTGTCA GAATAGGAAA ACATTGCAAT AAATGGCTTG TTAITITCAA AITICGACII CICCAACIIC GAIITITCCII IICCAACAAC AAITACI'IAI AGICCGAIAA IAAAIAACAI AATCCI'I'IA TAITAIAAAI gacaatctta agaaaataaa teeeggaaaa gacaeggtet gtaaegagag teaegaaaeg tacataateg agtgaettag aagtgetgtt acaaetette GITTGGITCI TCACCCACGG GAACCATCGI CGGACCCCAC TGGAGAICIC GACCTCCGAC ACCCTGAGGI CCCCGGGGGC ACAAGICCIG TGIAGAIAAC GTCTCTGAGT AAAGTGTCGG AAAGCAAGAC GACTGGTTTA CCGGTCAAAA GACCATCCTT CTACCTCCAA ATGGTCAACA AATCTTTGTC TTTATCTGAA aataaaggit taaagcigaa gaggiigaag ctaaaaggaa aaggiigiig ttaatgaata tcaggciatt atttattgta ttaggaaaat ataatattta TGTTCAGGAC ACATCTATTG AAATAGACTT CCGGTCCGAG AAGGAACTTA GAAGAGGAA CAGGACGAAC CCAAGTATCC TTAACCATTC CGGAGACCTG ACCGGACAGA CCGGGGACTC TCACCACGGG CCCACACCCCA 2001 GGCCAGGCTC TTCCTTGAAT CTTCTCCCTT GTCCTGCTTG GGTTCATAGG AATTGGTAAG GCCTCTGGAC TGGCCTGTCT GGCCCTGAG AGTGGTGCCC GGGTGTGGGT CAGAGACTCA TITCACAGCC TITCGITCTG CTGACCAAAI GGCCAGTTTT CTGGTAGGAA GATGGAGGTT TACCAGTTGT TTAGAAACAG TGGAACACTC CTCTACTCTT ACAGAGCCTT GAGAGACCCA GCTGCAGACC ATGCCAGACC CACTGAAATG ACCAAGACAG GTTCAGGTAG ACCITGIGAG GAGATGAGAA TGTCTCGGAA CICTCTGGGT CGACGICTGG TACGGTCTGG GTGACTTTAC TGGTTCTGTC CAAGICCATC CAAACCAAGA AGTGGGTGCC CTTGGTAGCA GCCTGGGGTG ACCTCTAGAG CTGGAGGCTG TGGGACTCCA GGGGCCCCCG CTGTTAGAAT TCTTTTATTT AGGGCCTTTT 2701 2401 2501 2201 2301 2101

14/27

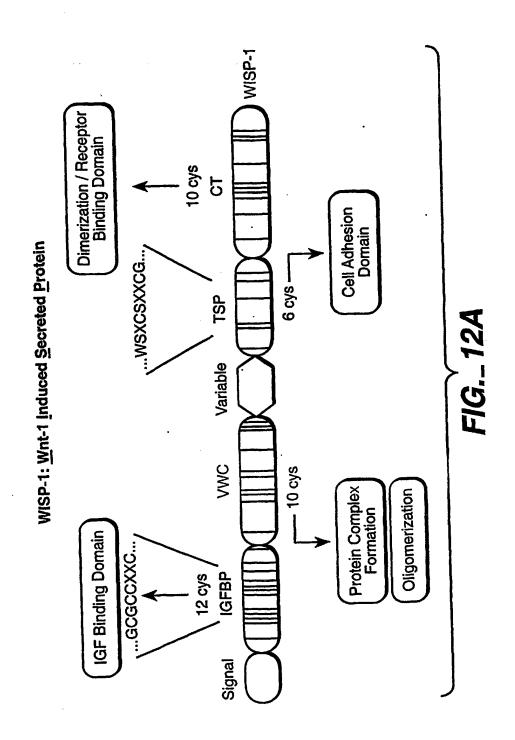
		Effect of WISP-1 Expression on NRK Metastatic Potential									
Weeks after Innoculation	Incidence of Lung Metastasis										
	NRK/Control	NRK/WISP-1L	NRK/WISP-1H								
2	. 0/1	0/1	2/2 (Grade I)								
3	0/4	3/3 (Grade I)	4/4 (Grade I-II)								
4	0/4	4/4 (Grade I)	3/3 (Grade III)								
2	0/2	0/2	1/2 (Grade I)								
	0/4	2/4 (Grade I)	3/4 (Grade I)								
4	0/4	4/4 (Grade I)	3/4 (Grade I-II)								
	Innoculation 2	NRK/Control	NRK/Control NRK/WISP-1L 2								

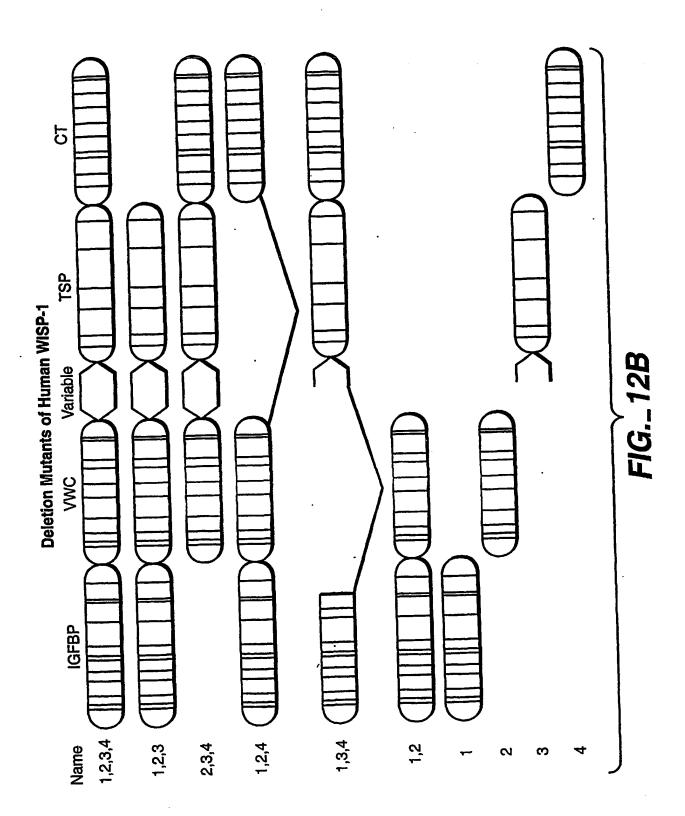
FIG._10



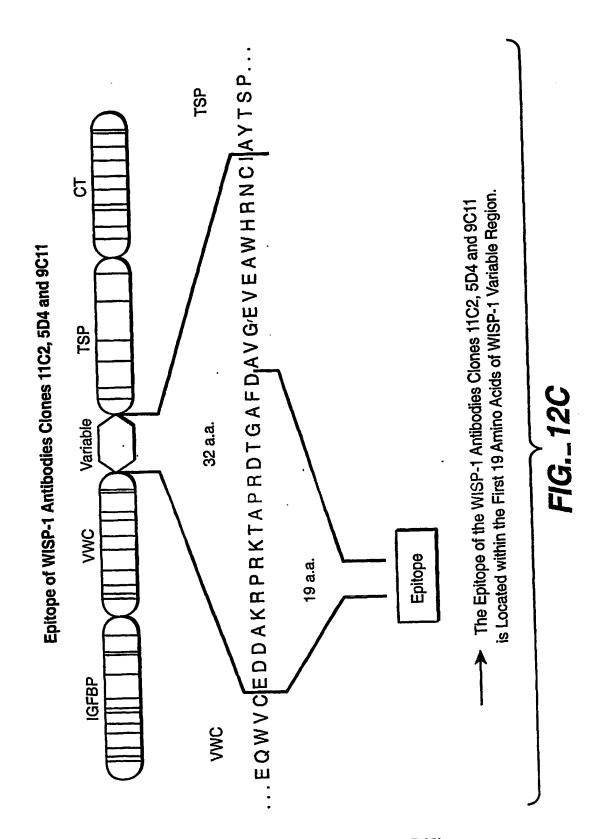
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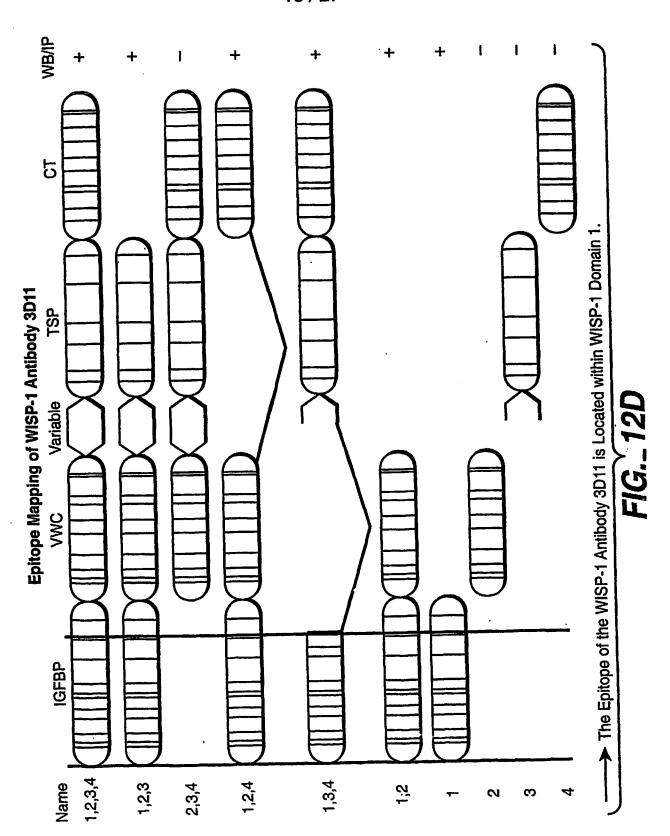
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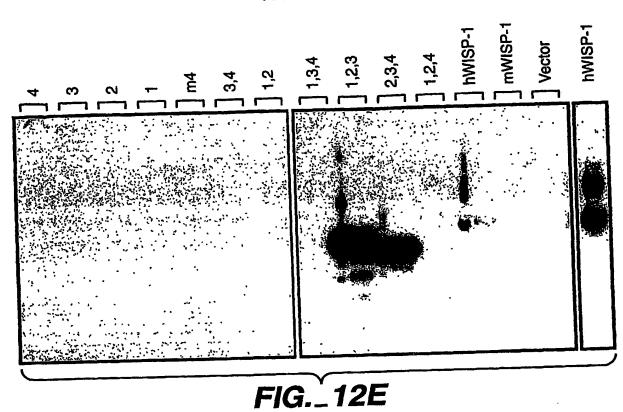


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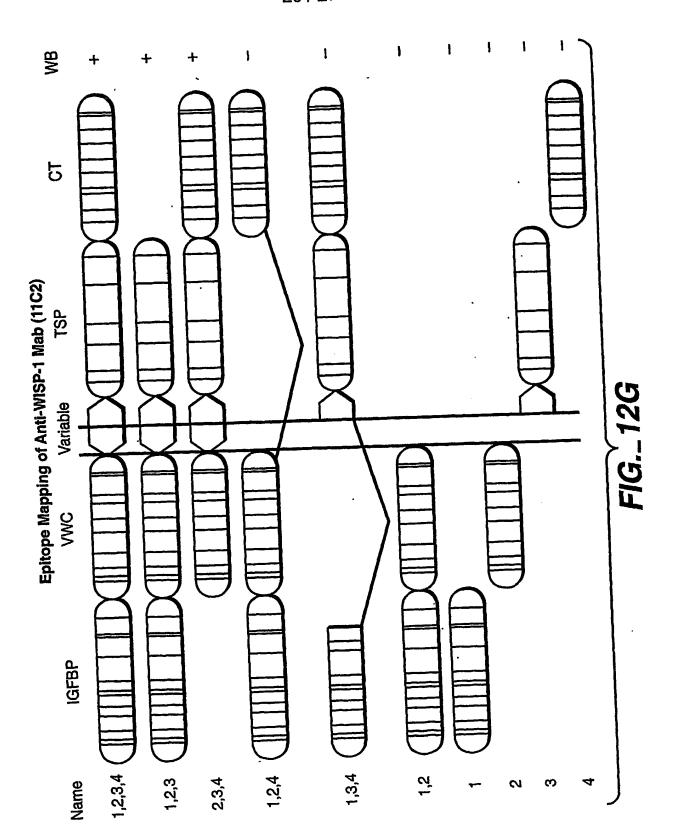


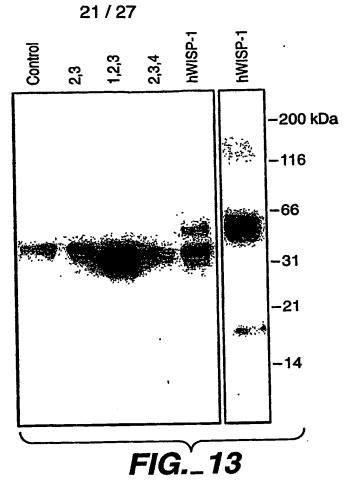
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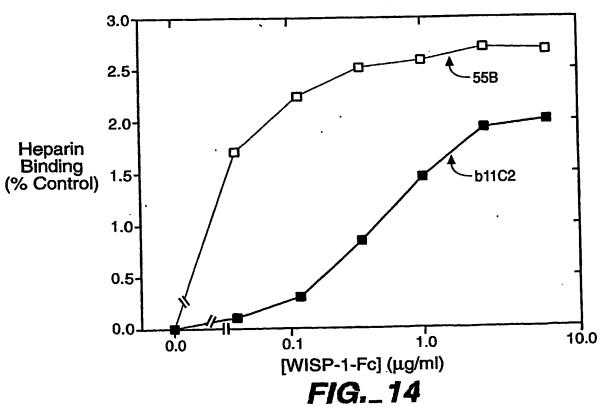


1,2 3,4 1,2,3 1,2,4 1,2,4 2,3,4 hwiSP-1

FIG._12F
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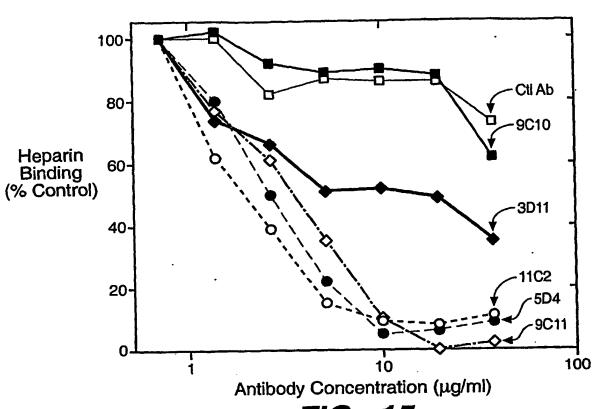


FIG._15

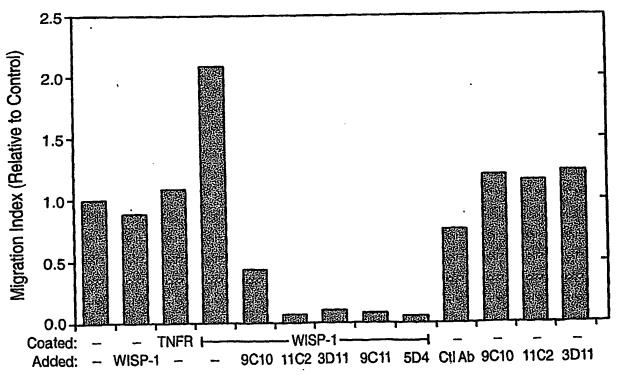


FIG._ 16A

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WISP-1 Antibodies Characterization											
Clones Isoty		Epitope			1400	Blocking Activity					
	Isotype	Struct.	Cross Compet.	ΙP	WB	Heparin Binding	Cell Migrat.	Lung Met.			
3D11	2b	Domain '1	Α	_	+	+/	+	+			
9C10	2b	Domain 1	В	+		-	+	+			
11C2	2b	Variable Region	С	+	+	+	+	+			
5D4	2a	Variable Region	C	+	+	+	+	+			
9C11	2a	Variable Region	С	+	+	+	+	+			

FIG._16B

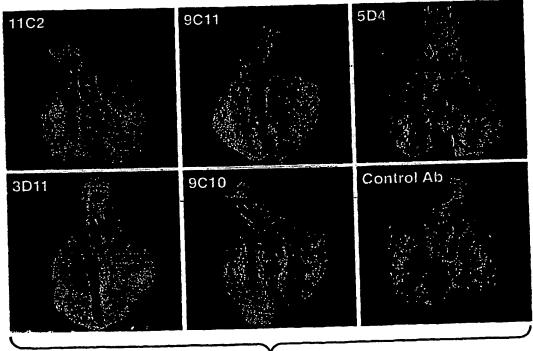


FIG._17A

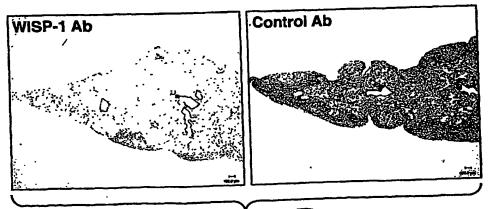
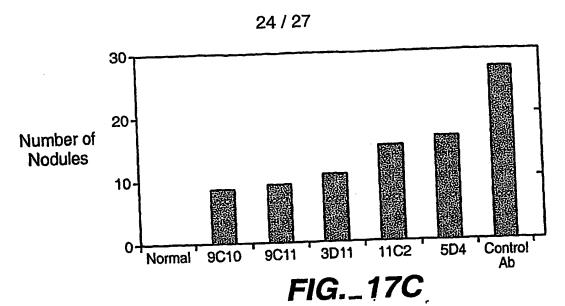
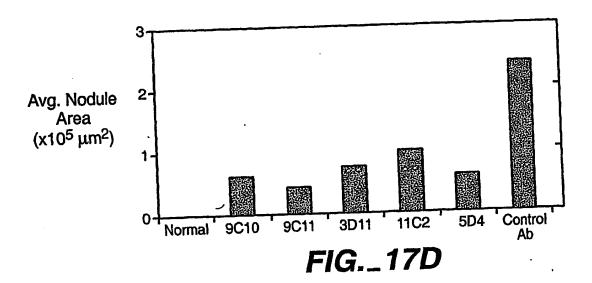
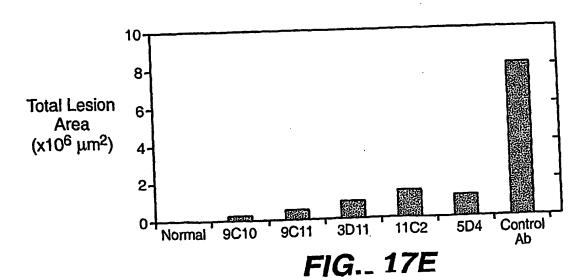


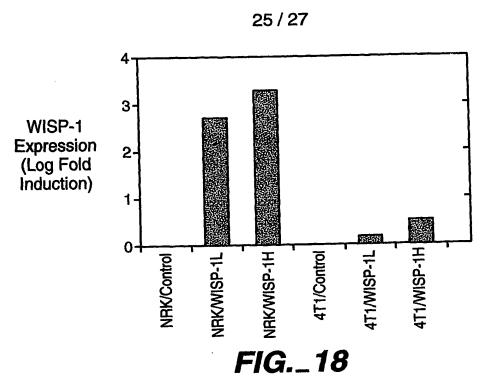
FIG._17B
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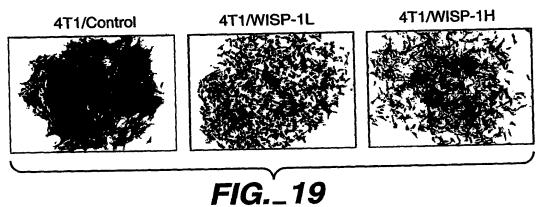






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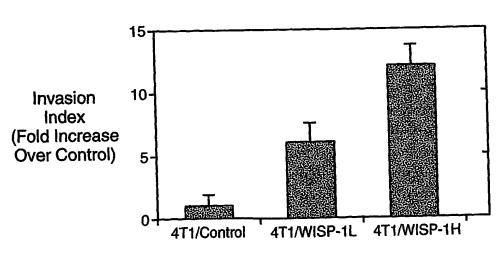
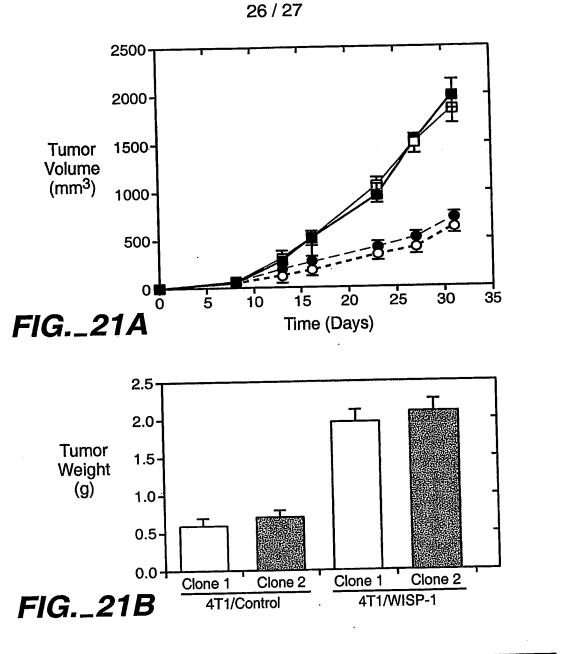


FIG._20
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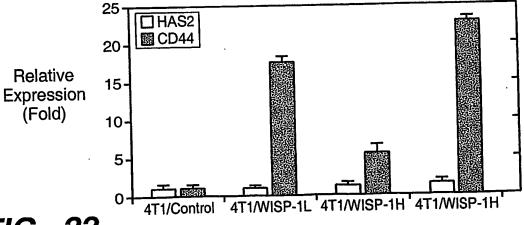
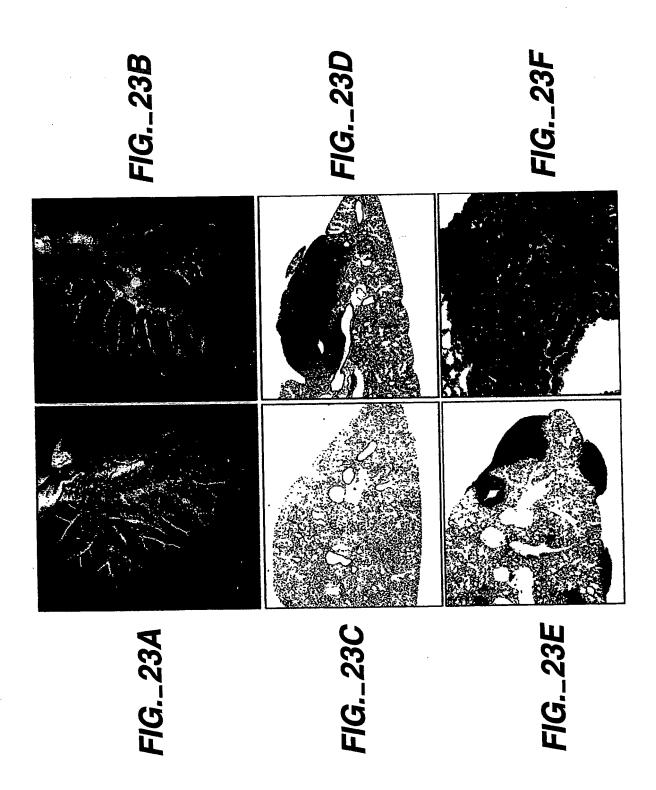


FIG._22



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